# INTERNATIONAL SEARCH REPORT

International application No. PCT/US00/03583

A. CL	ASSIFICATION OF SUBJECT MATTER			
IPC(7) :A61K 38/00; C07K 1/00; C07H 21/04; C12P 21/06; G01N 33/566				
US CL:514/2; 530/350, 388.22; 536/23.5; 435/69.1; 436/501 According to International Patent Classification (IPC) or to both national classification and IPC				
B. FIELDS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols)				
U.S. : 514/2; 530/350, 388.22; 536/23.5; 435/69.1; 436/501				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)				
Please See Extra Sheet.				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
The state of the s				
Category*	Citation of document, with indication, where	appropriate, of the relevant passages	Relevant to claim No.	
X	SPARATORE et al. Extracellular hig	1-3		
	essential for murine erythroleukemia c			
	August 1996, Vol. 320, pages 253-256, see the Abstract.			
A	WANG, H. et al. HMG-1 as a late me	1-8		
	mice. Science. 09 July 1999, Vol. 285 document.			
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	er documents are listed in the continuation of Box	٠		
*A* document defining the general state of the art which is not social and		"T" later document published after the inter date and not in conflict with the appli- the principle or theory underlying the	cation but cited to understand	
	ier document published on or after the international filing date	"X" document of particular relevance; the considered novel or cannot be considered	claimed invention cannot be	
cite	ument which may throw doubts on priority claim(s) or which is d to establish the publication date of another citation or other	when the document is taken alone	·	
	cial reason (as specified)  ument referring to an oral disclosure, use, exhibition or other uns	considered to involve an inventive combined with one or more other such	step when the document is documents, such combination	
"P" doc	ument published prior to the international filing date but later than priority date claimed	being obvious to a person skilled in the		
Date of the actual completion of the international search		Date of mailing of the international search report		
28 MARCH 2001		23 MAY 2001		
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks		Authorized officer age 3	uelces	
Box PCT	D.C. 20231	MICHAEL T. BRANNOCK	riogers .	
Facsimile No		Telephone No. (703) 308-0196	0	

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International application No. PCT/US00/03583

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)	_			
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:				
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:				
2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:				
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).				
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)				
This International Searching Authority found multiple inventions in this international application, as follows:				
Please See Extra Sheet.				
1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.				
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.				
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:				
4. X No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  1-8				
Remark on Protest				
No protest accompanied the payment of additional search fees.				

### INTERNATIONAL SEARCH REPORT

International application No. PCT/US00/03583

#### B. FIELDS SEARCHED

Electronic data bases consulted (Name of data base and where practicable terms used):

STN: MEDLINE, BIOSIS, BIOTECHNO; WEST 2.0

search terms: HMG1 antagonist, anti-HMG1, antibody, sepsis, inflammatory cascade, high mobility group, amphoterin, anti-amphoterin

### BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING

This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be searched, the appropriate additional search fees must be paid.

Group I, claim(s)1-8, drawn to a pharmaceutical composition for treating disease comprising an antagonist of HMG1 and a method for treating disease characterized by activation of the inflammatory cytokine cascade.

Group II, claim(s) 9-11, drawn to a diagnostic method.

Group III, claim(s) 12, drawn to a pharmaceutical composition for effecting weight loss.

Group IV, claim 13, drawn to a method for effecting weight loss.

This application contains claims directed to more than one species of the generic invention. These species are deemed to lack Unity of Invention because they are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for more than one species to be searched, the appropriate additional search fees must be paid. The species are as follows:

A pharmaceutical composition comprising:

- i) an antibody,
- ii) an antisense nucleic acid.
- iii) and HMG1 receptor antagonist

The claims are deemed to correspond to the species listed above in the following manner:

Claims 1-8 require the above species of compounds

The following claims are generic: 1-8

The inventions listed as Groups do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: Pursuant to 37 C.F.R. 1.475(d), this Authority considers that the main invention in the instant application comprises the first recited product, a pharmaceutical composition comprising and HMG1 receptor antagonist and the first recited method of using that product, namely in the process of treating a condition characterized by activation of the inflammatory cytokine cascade. Further, pursuant to 37 C.F.R. 1.475(b)-(d), the ISA/US considers that the materially and functionally dissimilar products of Groups III and the additional methods of Groups II and IV do not correspond to the main invention. This Authority therefore considers that the several inventions do not share a special technical feature within the meaning of PCT Rule 13.2 and thus do not relate to a single general inventive concept within the meaning of PCT Rule 13.1.

The species listed above do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, the species lack the same or corresponding special technical features for the following reasons: Antibodies, nucleic acids and molecules which function as HMG1 receptor antagonists are functionally and structurally distinct compounds.